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IN THE UNITED STATES PATENT AND TRADEMARK OFFICE



Applicants: Jordan J.N. Tang and Arun K. Ghosh

Serial No.: 09/506,988 Art Unit: 1625

Filed: February 18, 2000 Examiner: Seaman, D.

For: *PROTEASE INHIBITORS THAT OVERCOME DRUG RESISTANCE*

Assistant Commissioner for Patents
Washington, D.C. 20231

**SUPPLEMENTAL INFORMATION DISCLOSURE STATEMENT**

Sir:

Pursuant to the duty of disclosure under 37 C.F.R. §1.56 and 37 C.F.R. §1.97, Applicants submit a Supplemental Information Disclosure Statement, including two (2) pages of Form PTO-1449, copies of the documents cited therein, and a copy of the International Search Report, mailed 13 November 2000, in the corresponding PCT application PCT/US00/04215.

Enclosed is a check for $240.00 representing the fee required under 37 C.F.R. §1.17(p) for an Information Disclosure Statement filed after a first office action on the merits under 37 C.F.R. §1.97(c). It is believed that no additional fees are required with this submission. However, should a fee be required, the Commissioner is hereby authorized to charge any fees to Deposit Account No. 01-2507.

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02/28/2001 TGEDAMU1 0000102436 ME

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01 FC:126 180.00 OP

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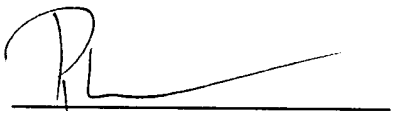
**Foreign Documents**

| Number | Publication Date | Patentee | Country |
|---|---|---|---|
| 0 373 497 A2 | 06-20-1990 | Hoechst Aktiengesellschaft | EP |
| WO 92/03472 A1 | 03-05-1992 | The Upjohn Company | PCT |

**Remarks**

This statement should not be interpreted as a representation that an exhaustive search has been conducted or that no better art exists. Moreover, Applicants invite the Examiner to make an independent evaluation of the cited art to determine its relevance to the subject matter of the present application. Applicants are of the opinion that their claims patentably distinguish over the art referred to herein, either alone or in combination.

Respectfully submitted,



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Dated: February 22, 2001

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**Publications**

BAKER & CONDON, "Dipeptide isosteres. 1. Synthesis of dihydroxyethylene dipeptide isosteres via diastereoselective additions of alkyllithium reagents to *N,N*-dimethylhydrazones. Preparations of renin and HIV-1 protease inhibitors transition-state mimics," *J Org Chem* 58:3277-3284 (1993).

BAKER, et al., "Nonpeptide renin inhibitors employing a novel 3-Aza (or oxa)-2,4-dialkyl glutaric acid moiety as a $P_2/P_3$ amide bond replacement," *J Med Chem* 35:1722-1734 (1992).

BENNETT, et al., "The synthesis of novel HIV-protease inhibitors via silica gel assisted addition of amines to epoxides," *SYNLETT* 9:703-704 (1993).

DREYER, et al., "Inhibition of human immunodeficiency virus 1 protease in vitro: rational design of design of substrate analogue inhibitors," *Proc Natl Acad Sci. USA* 86:9752-9756 (1989).

MARINIER, et al., "HIV-1 protease inhibitors: ketomethylene isosteres with unusually high affinity compared with hydroxyethylene isostere analogs," *Bioorganic & Medicinal Chemistry* 2(9):919-925 (1994).

**Certificate of Mailing under 37 CFR §1.8(a)**

I hereby certify that this Supplemental Information Disclosure Statement, along with any paper referred to as being attached or enclosed, is being deposited with the United States Postal Service on the date shown below with sufficient postage as first-class mail in an envelope addressed to the Assistant Commissioner for Patents, Washington, D.C. 20231.

Date: February 22, 2001

Kimberly L. Adams